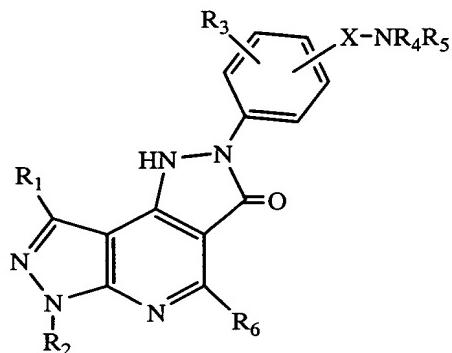


What is claimed is:

1. A compound of formula I



(I)

wherein

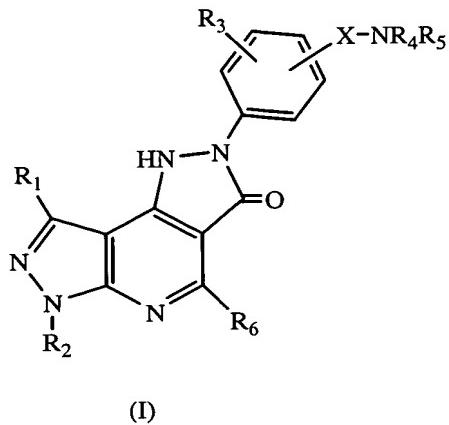
- 5        X is CO or SO<sub>2</sub>;
- R<sub>1</sub> and R<sub>2</sub> are each independently H, C<sub>1</sub>-C<sub>10</sub>alkyl optionally substituted with one or more halogen, hydroxy, C<sub>1</sub>-C<sub>4</sub>alkoxy, CO<sub>2</sub>R<sub>8</sub>, CONR<sub>9</sub>R<sub>10</sub>, C<sub>3</sub>-C<sub>7</sub>cycloalkyl or optionally substituted phenyl groups, or phenyl optionally substituted with one to three halogen, hydroxy, C<sub>1</sub>-C<sub>6</sub>haloalkyl, C<sub>1</sub>-C<sub>4</sub>alkoxy, CO<sub>2</sub>R<sub>11</sub>, NR<sub>12</sub>R<sub>13</sub> or CN groups;
- 10      R<sub>3</sub> is H, F, Cl, Br or I;
- R<sub>4</sub> and R<sub>5</sub> are each independently H, NH<sub>2</sub>, CH<sub>2</sub>CH<sub>2</sub>OCH<sub>2</sub>CH<sub>2</sub>OCH<sub>2</sub>CH<sub>2</sub>NH<sub>2</sub> or a C<sub>1</sub>-C<sub>6</sub>alkyl group optionally substituted with one or two CN, OR<sub>14</sub>, NR<sub>15</sub>R<sub>16</sub>, CO<sub>2</sub>R<sub>17</sub> or C<sub>3</sub>-C<sub>7</sub>cycloalkyl group,
- 15      phenyl optionally substituted with one or two halogen, CN, OR<sub>14</sub>, NR<sub>15</sub>R<sub>16</sub>, CO<sub>2</sub>R<sub>17</sub>, COR<sub>18</sub>, an optionally substituted C<sub>1</sub>-C<sub>6</sub>alkyl or an optionally substituted C<sub>2</sub>-C<sub>6</sub>alkenyl group,
- benzyl optionally substituted with one or two halogen, OR<sub>14</sub>, COR<sub>18</sub>, or a C<sub>1</sub>-C<sub>3</sub>alkyl group optionally substituted with one OR<sub>14</sub> group, or
- 20      pyridinyl optionally substituted with one or two halogen, OR<sub>14</sub>, NR<sub>15</sub>R<sub>16</sub> or CO<sub>2</sub>R<sub>17</sub> groups, or
- R<sub>4</sub> and R<sub>5</sub> may be taken together with the atom to which they are attached to form an optionally substituted 5- to 7-membered ring optionally

- containing one double bond, a benzofused ring or an additional heteroatom selected from O, NR<sub>19</sub> or S;
- R<sub>6</sub> is phenyl optionally substituted with one to three halogen, NO<sub>2</sub>, CN, hydroxy, C<sub>1</sub>-C<sub>6</sub>alkyl, C<sub>1</sub>-C<sub>6</sub>alkylthio, C<sub>1</sub>-C<sub>6</sub>haloalkyl, C<sub>1</sub>-C<sub>6</sub>alkoxy, phenyl, phenoxy, benzyl, benzyloxy, SO<sub>n</sub>R<sub>20</sub>, SO<sub>2</sub>NR<sub>21</sub>R<sub>22</sub>, CO<sub>2</sub>R<sub>23</sub> or NR<sub>24</sub>R<sub>25</sub> groups,
- cycloheteroalkyl optionally substituted with one or more halogen, NO<sub>2</sub>, CN, hydroxy, C<sub>1</sub>-C<sub>6</sub>alkyl, C<sub>1</sub>-C<sub>6</sub>alkylthio, C<sub>1</sub>-C<sub>6</sub>haloalkyl, C<sub>1</sub>-C<sub>6</sub>alkoxy, phenyl, phenoxy, benzyl, benzyloxy, SO<sub>n</sub>R<sub>20</sub>, SO<sub>2</sub>NR<sub>21</sub>R<sub>22</sub>, CO<sub>2</sub>R<sub>23</sub> or NR<sub>24</sub>R<sub>25</sub> groups, or
- heteroaryl optionally substituted with one or more halogen, NO<sub>2</sub>, CN, hydroxy, C<sub>1</sub>-C<sub>6</sub>alkyl, C<sub>1</sub>-C<sub>6</sub>alkylthio, C<sub>1</sub>-C<sub>6</sub>haloalkyl, C<sub>1</sub>-C<sub>6</sub>alkoxy, phenyl, phenoxy, benzyl, benzyloxy, SO<sub>n</sub>R<sub>20</sub>, SO<sub>2</sub>NR<sub>21</sub>R<sub>22</sub>, CO<sub>2</sub>R<sub>23</sub> or NR<sub>24</sub>R<sub>25</sub> groups;
- R<sub>8</sub>, R<sub>11</sub>, R<sub>17</sub>, R<sub>18</sub> and R<sub>23</sub> are each independently H or a C<sub>1</sub>-C<sub>6</sub>alkyl, C<sub>3</sub>-C<sub>7</sub>cycloalkyl, C<sub>1</sub>-C<sub>6</sub>haloalkyl, phenyl, C<sub>5</sub>-C<sub>7</sub>cycloheteroalkyl or heteroaryl group each optionally substituted;
- R<sub>9</sub>, R<sub>10</sub>, R<sub>12</sub>, R<sub>13</sub>, R<sub>15</sub>, R<sub>16</sub>, R<sub>21</sub>, R<sub>22</sub>, R<sub>24</sub> and R<sub>25</sub> are each independently H or a C<sub>1</sub>-C<sub>6</sub>alkyl, C<sub>3</sub>-C<sub>7</sub>cycloalkyl, C<sub>1</sub>-C<sub>6</sub>haloalkyl, phenyl, C<sub>5</sub>-C<sub>7</sub>cycloheteroalkyl or heteroaryl group each optionally substituted or each of R<sub>9</sub> and R<sub>10</sub> or R<sub>12</sub> and R<sub>13</sub> or R<sub>15</sub> and R<sub>16</sub> or R<sub>21</sub> and R<sub>22</sub> or R<sub>24</sub> and R<sub>25</sub> may be taken together with the nitrogen atom to which they are attached to form a 5- to 7-membered ring optionally containing another heteroatom selected from O, N or S;
- n is 0 or an integer of 1 or 2;
- R<sub>14</sub> is H, C<sub>1</sub>-C<sub>3</sub>alkyl or C<sub>1</sub>-C<sub>3</sub>haloalkyl;
- R<sub>19</sub> is H or C<sub>1</sub>-C<sub>3</sub>alkyl; and
- R<sub>20</sub> is a C<sub>1</sub>-C<sub>6</sub>alkyl, C<sub>3</sub>-C<sub>7</sub>cycloalkyl, C<sub>1</sub>-C<sub>6</sub>haloalkyl, phenyl, C<sub>5</sub>-C<sub>7</sub>cyclo-heteroalkyl or heteroaryl group each optionally substituted; or
- the stereoisomers thereof or the pharmaceutically acceptable salts thereof.

2. The compound according to claim 1 wherein X is CO.

3. The compound according to claim 1 wherein R<sub>1</sub> is H.
4. The compound according to claim 1 wherein R<sub>1</sub> is H. The compound according to claim 1 wherein R<sub>6</sub> is a phenyl group optionally substituted with one or two CN, NO<sub>2</sub>, halogen, CF<sub>3</sub>, C<sub>1</sub>-C<sub>3</sub>alkoxy or CO<sub>2</sub>R<sub>23</sub> groups.
- 5 5. The compound according to claim 2 wherein R<sub>2</sub> is H or C<sub>1</sub>-C<sub>3</sub>alkyl.
6. The compound according to claim 2 wherein R<sub>4</sub> and R<sub>5</sub> are each independently H or a C<sub>1</sub>-C<sub>3</sub>alkyl, phenyl or benzyl group each optionally substituted with one or two hydroxy groups or R<sub>4</sub> and R<sub>5</sub> may be taken together with the atom to which they are attached to form a pyrrolidinyl or morpholinyl ring each optionally substituted with one carboxy group.
- 10 7. The compound according to claim 5 wherein R<sub>6</sub> is phenyl optionally substituted in the 3-position with CF<sub>3</sub>.
8. The compound according to claim 7 wherein R<sub>1</sub> is H.
9. The compound according to claim 7 wherein R<sub>1</sub> is H. The compound according to claim 1 selected from the group consisting of:
- 15 N-(4-hydroxyphenyl)-3-(6-methyl-3-oxo-4-[3-(trifluoromethyl)phenyl]-3,6-dihydrodipyrazolo[3,4-b:3,4-d]pyridin-2(1H)-yl)benzamide;
- N-(2,2-dimethoxyethyl)-N-methyl-3-(6-methyl-3-oxo-4-[3-(trifluoromethyl)phenyl]-3,6-dihydrodipyrazolo[3,4-b:3,4-d]pyridin-2(1H)-yl)benzamide;
- 20 6-methyl-2-[3-(1-pyrrolidinylcarbonyl)phenyl]-4-[3-(trifluoromethyl)phenyl]-1,6-dihydrodipyrazolo[3,4-b:3,4-d]pyridin-3(2H)-one;
- (2R)-1-[3-(6-methyl-3-oxo-4-[3-(trifluoromethyl)phenyl]-3,6-dihydrodipyrazolo[3,4-b:3,4-d]pyridin-2(1H)-yl)benzoyl]-2-pyrrolidinecarboxylic acid;
- N-(3,4-dihydroxybenzyl)-3-(6-methyl-3-oxo-4-[3-(trifluoromethyl)phenyl]-3,6-dihydrodipyrazolo[3,4-b:3,4-d]pyridin-2(1H)-yl)benzamide;
- 25 N-(2-hydroxypropyl)-3-(6-methyl-3-oxo-4-[3-(trifluoromethyl)phenyl]-3,6-dihydrodipyrazolo[3,4-b:3,4-d]pyridin-2(1H)-yl)benzamide;
- 1-{2-chloro-5-[6-methyl-3-oxo-4-[3-(trifluoromethyl)phenyl]-3,6-dihydrodipyrazolo[3,4-b:3',4'-d]pyridin-2(1H)-yl]benzoyl}-D-proline;

- 2-(4-chloro-3-{{(2R)-2-(hydroxymethyl)pyrrolidin-1-yl}carbonyl}phenyl)-6-methyl-4-[3-(trifluoromethyl)phenyl]-1,6-dihydrodipyrazolo[3,4-b:3,4-d]pyridin-3(2H)-one; N-(4-hydroxyphenyl)-4-(6-methyl-3-oxo-4-[3-(trifluoromethyl)phenyl]-3,6-dihydrodipyrazolo[3,4-b:3',4'-d]pyridin-2(1H)-yl)benzamide;
- 5 N-(2-hydroxyphenyl)-4-(6-methyl-3-oxo-3-[3-(trifluoromethyl)phenyl]-3,6-dihydrodipyrazolo[3,4-b:3',4'-d]pyridin-2(1H)-yl)benzamide; 6-methyl-2-[4-(4-morpholinylcarbonyl)phenyl]-4-[3-(trifluoromethyl)phenyl]-1,6-dihydrodipyrazolo[3,4-b:3,4-d]pyridin-3(2H)-one; N-[4-(2-hydroxyethyl)phenyl]-4-(6-methyl-3-oxo-4-[3-(trifluoromethyl)phenyl]-3,6-
- 10 dihydrodipyrazolo[3,4-b:3,4-d]pyridin-2(1H)-yl)benzamide; N-[3-(1-hydroxyethyl)phenyl]-4-(6-methyl-3-oxo-4-[3-(trifluoromethyl)phenyl]-3,6-dihydrodipyrazolo[3,4-b:3,4-d]pyridin-2(1H)-yl)benzamide; N-[3-(hydroxymethyl)phenyl]-4-(6-methyl-3-oxo-4-[3-(trifluoromethyl)phenyl]-3,6-dihydrodipyrazolo[3,4-b:3,4-d]pyridin-2(1H)-yl)benzamide;
- 15 N-(5-hydroxpentyl)-4-(6-methyl-3-oxo-4-[3-(trifluoromethyl)phenyl]-3,6-dihydrodipyrazolo[3,4-b:3,4-d]pyridin-2(1H)-yl)benzenesulfonamide; N-benzyl-4-[6-methyl-3-oxo-4-(3-trifluoromethyl-phenyl)-3,6-dihydro-1H-1,2,5,6,7-pentaaza-as-indacen-2-yl]-benzenesulfonamide; N-(2-hydroxyethyl)-4-(6-methyl-3-oxo-4-[3-(trifluoromethyl)phenyl]-3,6-
- 20 dihydrodipyrazolo[3,4-b:3,4-d]pyridin-2(1H)-yl)benzenesulfonamide; methyl ({[4-(6-methyl-3-oxo-4-[3-(trifluoromethyl)phenyl]-3,6-dihydrodipyrazolo[3,4-b:3,4-d]pyridin-2(1H)-yl)phenyl]sulfonyl}amino)acetate; N-cyclopropylmethyl-4-[6-methyl-3-oxo-4-(3-trifluoromethyl-phenyl)-3,6-dihydro-1H-1,2,5,6,7-pentaaza-as-indacen-2-yl]-benzenesulfonamide;
- 25 ({[4-(6-methyl-3-oxo-4-[3-(trifluoromethyl)phenyl]-3,6-dihydrodipyrazolo[3,4-b:3,4-d]pyridin-2(1H)-yl)phenyl]sulfonyl}amino)acetic acid; the stereoisomers thereof; and the pharmaceutically acceptable salts thereof.
- 30 10. A method for the treatment of an immune disorder related to or affected by the immune regulatory protein B7-1 which comprises providing a patient in need thereof an immunotherapeutically effective amount of a compound of formula I



wherein

**X** is CO or SO<sub>2</sub>;

**R**<sub>1</sub> and **R**<sub>2</sub> are each independently H, C<sub>1</sub>-C<sub>10</sub>alkyl optionally substituted with one or more halogen, hydroxy, C<sub>1</sub>-C<sub>4</sub>alkoxy, CO<sub>2</sub>R<sub>8</sub>, CONR<sub>9</sub>R<sub>10</sub>, C<sub>3</sub>-C<sub>7</sub>cycloalkyl or optionally substituted phenyl groups, or phenyl optionally substituted with one to three halogen, hydroxy, C<sub>1</sub>-C<sub>6</sub>haloalkyl, C<sub>1</sub>-C<sub>4</sub>alkoxy, CO<sub>2</sub>R<sub>11</sub>, NR<sub>12</sub>R<sub>13</sub> or CN groups;

**R**<sub>3</sub> is H, F, Cl, Br or I;

**R**<sub>4</sub> and **R**<sub>5</sub> are each independently H, NH<sub>2</sub>, CH<sub>2</sub>CH<sub>2</sub>OCH<sub>2</sub>CH<sub>2</sub>OCH<sub>2</sub>CH<sub>2</sub>NH<sub>2</sub> or a C<sub>1</sub>-C<sub>6</sub>alkyl group optionally substituted with one or two CN, OR<sub>14</sub>, NR<sub>15</sub>R<sub>16</sub>, CO<sub>2</sub>R<sub>17</sub> or C<sub>3</sub>-C<sub>7</sub>cycloalkyl group, phenyl optionally substituted with one or two halogen, CN, OR<sub>14</sub>, NR<sub>15</sub>R<sub>16</sub>, CO<sub>2</sub>R<sub>17</sub>, COR<sub>18</sub>, an optionally substituted C<sub>1</sub>-C<sub>6</sub>alkyl or an optionally substituted C<sub>2</sub>-C<sub>6</sub>alkenyl group, benzyl optionally substituted with one or two halogen, OR<sub>14</sub>, COR<sub>18</sub>, or a C<sub>1</sub>-C<sub>3</sub>alkyl group optionally substituted with one OR<sub>14</sub> group, or pyridinyl optionally substituted with one or two halogen, OR<sub>14</sub>, NR<sub>15</sub>R<sub>16</sub> or CO<sub>2</sub>R<sub>17</sub> groups, or

**R**<sub>4</sub> and **R**<sub>5</sub> may be taken together with the atom to which they are attached to form an optionally substituted 5- to 7-membered ring optionally containing one double bond, a benzofused ring or an additional heteroatom selected from O, NR<sub>19</sub> or S;

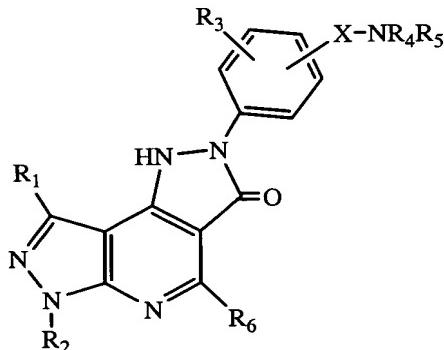
- R<sub>6</sub> is phenyl optionally substituted with one to three halogen, NO<sub>2</sub>, CN, hydroxy, C<sub>1</sub>-C<sub>6</sub>alkyl, C<sub>1</sub>-C<sub>6</sub>alkylthio, C<sub>1</sub>-C<sub>6</sub>haloalkyl, C<sub>1</sub>-C<sub>6</sub>alkoxy, phenyl, phenoxy, benzyl, benzyloxy, SO<sub>n</sub>R<sub>20</sub>, SO<sub>2</sub>NR<sub>21</sub>R<sub>22</sub>, CO<sub>2</sub>R<sub>23</sub> or NR<sub>24</sub>R<sub>25</sub> groups,
- 5        cycloheteroalkyl optionally substituted with one or more halogen, NO<sub>2</sub>, CN, hydroxy, C<sub>1</sub>-C<sub>6</sub>alkyl, C<sub>1</sub>-C<sub>6</sub>alkylthio, C<sub>1</sub>-C<sub>6</sub>haloalkyl, C<sub>1</sub>-C<sub>6</sub>alkoxy, phenyl, phenoxy, benzyl, benzyloxy, SO<sub>n</sub>R<sub>20</sub>, SO<sub>2</sub>NR<sub>21</sub>R<sub>22</sub>, CO<sub>2</sub>R<sub>23</sub> or NR<sub>24</sub>R<sub>25</sub> groups, or
- 10      heteroaryl optionally substituted with one or more halogen, NO<sub>2</sub>, CN, hydroxy, C<sub>1</sub>-C<sub>6</sub>alkyl, C<sub>1</sub>-C<sub>6</sub>alkylthio, C<sub>1</sub>-C<sub>6</sub>haloalkyl, C<sub>1</sub>-C<sub>6</sub>alkoxy, phenyl, phenoxy, benzyl, benzyloxy, SO<sub>n</sub>R<sub>20</sub>, SO<sub>2</sub>NR<sub>21</sub>R<sub>22</sub>, CO<sub>2</sub>R<sub>23</sub> or NR<sub>24</sub>R<sub>25</sub> groups;
- 15      R<sub>8</sub>, R<sub>11</sub>, R<sub>17</sub>, R<sub>18</sub> and R<sub>23</sub> are each independently H or a C<sub>1</sub>-C<sub>6</sub>alkyl, C<sub>3</sub>-C<sub>7</sub> cycloalkyl, C<sub>1</sub>-C<sub>6</sub>haloalkyl, phenyl, C<sub>5</sub>-C<sub>7</sub>cycloheteroalkyl or heteroaryl group each optionally substituted;
- 20      R<sub>9</sub>, R<sub>10</sub>, R<sub>12</sub>, R<sub>13</sub>, R<sub>15</sub>, R<sub>16</sub>, R<sub>21</sub>, R<sub>22</sub>, R<sub>24</sub> and R<sub>25</sub> are each independently H or a C<sub>1</sub>-C<sub>6</sub>alkyl, C<sub>3</sub>-C<sub>7</sub>cycloalkyl, C<sub>1</sub>-C<sub>6</sub>haloalkyl, phenyl, C<sub>5</sub>-C<sub>7</sub>cycloheteroalkyl or heteroaryl group each optionally substituted or each of R<sub>9</sub> and R<sub>10</sub> or R<sub>12</sub> and R<sub>13</sub> or R<sub>15</sub> and R<sub>16</sub> or R<sub>21</sub> and R<sub>22</sub> or R<sub>24</sub> and R<sub>25</sub> may be taken together with the nitrogen atom to which they are attached to form a 5- to 7-membered ring optionally containing another heteroatom selected from O, N or S;
- 25      n is 0 or an integer of 1 or 2;
- R<sub>14</sub> is H, C<sub>1</sub>-C<sub>3</sub>alkyl or C<sub>1</sub>-C<sub>3</sub>haloalkyl;
- R<sub>19</sub> is H or C<sub>1</sub>-C<sub>3</sub>alkyl; and
- R<sub>20</sub> is a C<sub>1</sub>-C<sub>6</sub>alkyl, C<sub>3</sub>-C<sub>7</sub>cycloalkyl, C<sub>1</sub>-C<sub>6</sub>haloalkyl, phenyl, C<sub>5</sub>-C<sub>7</sub>cyclo-heteroalkyl or heteroaryl group each optionally substituted; or the stereoisomers thereof or the pharmaceutically acceptable salts thereof.
- 30      11.     The method according to claim 10 wherein said disorder is transplant rejection.

12. The method according to claim 10 wherein said disorder is an autoimmune disease.

13. The method according to claim 10 wherein said disorder is graft vs. 5 host disease.

14. The method according to claim 12 wherein said disease is multiple sclerosis or rheumatoid arthritis.

10 15. A pharmaceutical composition which comprises a pharmaceutically acceptable carrier and an effective amount of a compound of formula I



(I)

wherein

15 X is CO or SO<sub>2</sub>;

R<sub>1</sub> and R<sub>2</sub> are each independently H, C<sub>1</sub>-C<sub>10</sub>alkyl optionally substituted with one or more halogen, hydroxy, C<sub>1</sub>-C<sub>4</sub>alkoxy, CO<sub>2</sub>R<sub>8</sub>, CONR<sub>9</sub>R<sub>10</sub>, C<sub>3</sub>-C<sub>7</sub>cycloalkyl or optionally substituted phenyl groups, or phenyl optionally substituted with one to three halogen, hydroxy, C<sub>1</sub>-C<sub>6</sub>haloalkyl, C<sub>1</sub>-C<sub>4</sub>alkoxy, CO<sub>2</sub>R<sub>11</sub>, NR<sub>12</sub>R<sub>13</sub> or CN groups;

20 R<sub>3</sub> is H, F, Cl, Br or I;  
R<sub>4</sub> and R<sub>5</sub> are each independently H, NH<sub>2</sub>, CH<sub>2</sub>CH<sub>2</sub>OCH<sub>2</sub>CH<sub>2</sub>OCH<sub>2</sub>CH<sub>2</sub>NH<sub>2</sub> or a C<sub>1</sub>-C<sub>6</sub>alkyl group optionally substituted with one or two CN, OR<sub>14</sub>, NR<sub>15</sub>R<sub>16</sub>, CO<sub>2</sub>R<sub>17</sub> or C<sub>3</sub>-C<sub>7</sub>cycloalkyl group,

phenyl optionally substituted with one or two halogen, CN, OR<sub>14</sub>, NR<sub>15</sub>R<sub>16</sub>, CO<sub>2</sub>R<sub>17</sub>, COR<sub>18</sub>, an optionally substituted C<sub>1</sub>-C<sub>6</sub>alkyl or an optionally substituted C<sub>2</sub>-C<sub>6</sub>alkenyl group,

benzyl optionally substituted with one or two halogen, OR<sub>14</sub>, COR<sub>18</sub>, or a C<sub>1</sub>-C<sub>3</sub>alkyl group optionally substituted with one OR<sub>14</sub> group, or

pyridinyl optionally substituted with one or two halogen, OR<sub>14</sub>, NR<sub>15</sub>R<sub>16</sub> or CO<sub>2</sub>R<sub>17</sub> groups, or

R<sub>4</sub> and R<sub>5</sub> may be taken together with the atom to which they are attached to form an optionally substituted 5- to 7-membered ring optionally containing one double bond, a benzofused ring or an additional heteroatom selected from O, NR<sub>19</sub> or S;

R<sub>6</sub> is phenyl optionally substituted with one to three halogen, NO<sub>2</sub>, CN, hydroxy, C<sub>1</sub>-C<sub>6</sub>alkyl, C<sub>1</sub>-C<sub>6</sub>alkylthio, C<sub>1</sub>-C<sub>6</sub>haloalkyl, C<sub>1</sub>-C<sub>6</sub>alkoxy, phenyl, phenoxy, benzyl, benzyloxy, SO<sub>n</sub>R<sub>20</sub>, SO<sub>2</sub>NR<sub>21</sub>R<sub>22</sub>, CO<sub>2</sub>R<sub>23</sub> or NR<sub>24</sub>R<sub>25</sub> groups,

cycloheteroalkyl optionally substituted with one or more halogen, NO<sub>2</sub>, CN, hydroxy, C<sub>1</sub>-C<sub>6</sub>alkyl, C<sub>1</sub>-C<sub>6</sub>alkylthio, C<sub>1</sub>-C<sub>6</sub>haloalkyl, C<sub>1</sub>-C<sub>6</sub>alkoxy, phenyl, phenoxy, benzyl, benzyloxy, SO<sub>n</sub>R<sub>20</sub>, SO<sub>2</sub>NR<sub>21</sub>R<sub>22</sub>, CO<sub>2</sub>R<sub>23</sub> or NR<sub>24</sub>R<sub>25</sub> groups, or

heteroaryl optionally substituted with one or more halogen, NO<sub>2</sub>, CN, hydroxy, C<sub>1</sub>-C<sub>6</sub>alkyl, C<sub>1</sub>-C<sub>6</sub>alkylthio, C<sub>1</sub>-C<sub>6</sub>haloalkyl, C<sub>1</sub>-C<sub>6</sub>alkoxy, phenyl, phenoxy, benzyl, benzyloxy, SO<sub>n</sub>R<sub>20</sub>, SO<sub>2</sub>NR<sub>21</sub>R<sub>22</sub>, CO<sub>2</sub>R<sub>23</sub> or NR<sub>24</sub>R<sub>25</sub> groups;

R<sub>8</sub>, R<sub>11</sub>, R<sub>17</sub>, R<sub>18</sub> and R<sub>23</sub> are each independently H or a C<sub>1</sub>-C<sub>6</sub>alkyl, C<sub>3</sub>-C<sub>7</sub> cycloalkyl, C<sub>1</sub>-C<sub>6</sub>haloalkyl, phenyl, C<sub>5</sub>-C<sub>7</sub>cycloheteroalkyl or heteroaryl group each optionally substituted;

R<sub>9</sub>, R<sub>10</sub>, R<sub>12</sub>, R<sub>13</sub>, R<sub>15</sub>, R<sub>16</sub>, R<sub>21</sub>, R<sub>22</sub>, R<sub>24</sub> and R<sub>25</sub> are each independently H or a C<sub>1</sub>-C<sub>6</sub>alkyl, C<sub>3</sub>-C<sub>7</sub>cycloalkyl, C<sub>1</sub>-C<sub>6</sub>haloalkyl, phenyl, C<sub>5</sub>-C<sub>7</sub>cycloheteroalkyl or heteroaryl group each optionally substituted or each of R<sub>9</sub> and R<sub>10</sub> or R<sub>12</sub> and R<sub>13</sub> or R<sub>15</sub> and R<sub>16</sub> or R<sub>21</sub> and R<sub>22</sub> or R<sub>24</sub> and R<sub>25</sub> may be taken together with the nitrogen atom to which they are attached to form a 5- to 7-membered ring optionally containing another heteroatom selected from O, N or S;

n is 0 or an integer of 1 or 2;  
 R<sub>14</sub> is H, C<sub>1</sub>-C<sub>3</sub>alkyl or C<sub>1</sub>-C<sub>3</sub>haloalkyl;  
 R<sub>19</sub> is H or C<sub>1</sub>-C<sub>3</sub>alkyl; and  
 R<sub>20</sub> is a C<sub>1</sub>-C<sub>6</sub>alkyl, C<sub>3</sub>-C<sub>7</sub>cycloalkyl, C<sub>1</sub>-C<sub>6</sub>haloalkyl, phenyl, C<sub>5</sub>-C<sub>7</sub>cyclo-  
 5 heteroalkyl or heteroaryl group each optionally substituted; or  
 the stereoisomers thereof or the pharmaceutically acceptable salts thereof.

16. The composition according to claim 15 having a formula I compound  
 wherein X is CO.

10

17. The composition according to claim 16 having a formula I compound  
 wherein R<sub>1</sub> is H.

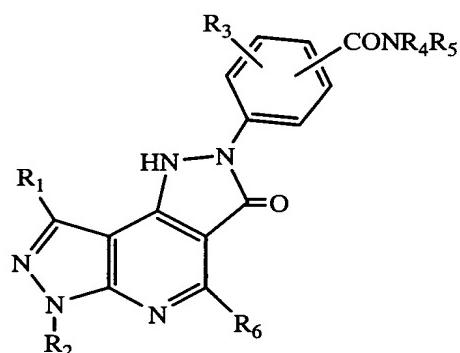
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18. The composition according to claim 17 having a formula I compound  
 wherein R<sub>2</sub> is H or CH<sub>3</sub>.

19. The composition according to claim 18 having a formula I compound  
 wherein R<sub>6</sub> is phenyl optionally substituted in the 3-position with CF<sub>3</sub>.

20

20. A process for the preparation of a compound of formula Ia



(Ia)

wherein

- R<sub>1</sub> and R<sub>2</sub> are each independently H, C<sub>1</sub>-C<sub>10</sub>alkyl optionally substituted with one or more halogen, hydroxy, C<sub>1</sub>-C<sub>4</sub>alkoxy, CO<sub>2</sub>R<sub>8</sub>, CONR<sub>9</sub>R<sub>10</sub>, C<sub>3</sub>-C<sub>7</sub>cycloalkyl or optionally substituted phenyl groups, or
- 5 phenyl optionally substituted with one to three halogen, hydroxy, C<sub>1</sub>-C<sub>6</sub>haloalkyl, C<sub>1</sub>-C<sub>4</sub>alkoxy, CO<sub>2</sub>R<sub>11</sub>, NR<sub>12</sub>R<sub>13</sub> or CN groups;
- R<sub>3</sub> is H, F, Cl, Br or I;
- R<sub>4</sub> and R<sub>5</sub> are each independently H, NH<sub>2</sub>, CH<sub>2</sub>CH<sub>2</sub>OCH<sub>2</sub>CH<sub>2</sub>OCH<sub>2</sub>CH<sub>2</sub>NH<sub>2</sub> or a
- 10 C<sub>1</sub>-C<sub>6</sub>alkyl group optionally substituted with one or two CN, OR<sub>14</sub>, NR<sub>15</sub>R<sub>16</sub>, CO<sub>2</sub>R<sub>17</sub> or C<sub>3</sub>-C<sub>7</sub>cycloalkyl group, phenyl optionally substituted with one or two halogen, CN, OR<sub>14</sub>, NR<sub>15</sub>R<sub>16</sub>, CO<sub>2</sub>R<sub>17</sub>, COR<sub>18</sub>, an optionally substituted C<sub>1</sub>-C<sub>6</sub>alkyl or an optionally substituted C<sub>2</sub>-C<sub>6</sub>alkenyl group,
- 15 benzyl optionally substituted with one or two halogen, OR<sub>14</sub>, COR<sub>18</sub>, or a C<sub>1</sub>-C<sub>3</sub>alkyl group optionally substituted with one OR<sub>14</sub> group, or pyridinyl optionally substituted with one or two halogen, OR<sub>14</sub>, NR<sub>15</sub>R<sub>16</sub> or CO<sub>2</sub>R<sub>17</sub> groups, or
- R<sub>4</sub> and R<sub>5</sub> may be taken together with the atom to which they are attached to form an optionally substituted 5- to 7-membered ring optionally containing one double bond, a benzofused ring or an additional heteroatom selected from O, NR<sub>19</sub> or S;
- 20 R<sub>6</sub> is phenyl optionally substituted with one to three halogen, NO<sub>2</sub>, CN, hydroxy, C<sub>1</sub>-C<sub>6</sub>alkyl, C<sub>1</sub>-C<sub>6</sub>alkylthio, C<sub>1</sub>-C<sub>6</sub>haloalkyl, C<sub>1</sub>-C<sub>6</sub>alkoxy, phenyl, phenoxy, benzyl, benzyloxy, SO<sub>n</sub>R<sub>20</sub>, SO<sub>2</sub>NR<sub>21</sub>R<sub>22</sub>, CO<sub>2</sub>R<sub>23</sub> or NR<sub>24</sub>R<sub>25</sub> groups,
- 25 cycloheteroalkyl optionally substituted with one or more halogen, NO<sub>2</sub>, CN, hydroxy, C<sub>1</sub>-C<sub>6</sub>alkyl, C<sub>1</sub>-C<sub>6</sub>alkylthio, C<sub>1</sub>-C<sub>6</sub>haloalkyl, C<sub>1</sub>-C<sub>6</sub>alkoxy, phenyl, phenoxy, benzyl, benzyloxy, SO<sub>n</sub>R<sub>20</sub>, SO<sub>2</sub>NR<sub>21</sub>R<sub>22</sub>, CO<sub>2</sub>R<sub>23</sub> or NR<sub>24</sub>R<sub>25</sub> groups, or
- 30 heteroaryl optionally substituted with one or more halogen, NO<sub>2</sub>, CN, hydroxy, C<sub>1</sub>-C<sub>6</sub>alkyl, C<sub>1</sub>-C<sub>6</sub>alkylthio, C<sub>1</sub>-C<sub>6</sub>haloalkyl, C<sub>1</sub>-C<sub>6</sub>alkoxy, phenyl, phenoxy, benzyl, benzyloxy, SO<sub>n</sub>R<sub>20</sub>, SO<sub>2</sub>NR<sub>21</sub>R<sub>22</sub>, CO<sub>2</sub>R<sub>23</sub> or NR<sub>24</sub>R<sub>25</sub> groups;

$R_8$ ,  $R_{11}$ ,  $R_{17}$ ,  $R_{18}$  and  $R_{23}$  are each independently H or a  $C_1$ - $C_6$ alkyl,  $C_3$ - $C_7$ cycloalkyl,  $C_1$ - $C_6$ haloalkyl, phenyl,  $C_5$ - $C_7$ cycloheteroalkyl or heteroaryl group each optionally substituted;

5         $R_9$ ,  $R_{10}$ ,  $R_{12}$ ,  $R_{13}$ ,  $R_{15}$ ,  $R_{16}$ ,  $R_{21}$ ,  $R_{22}$ ,  $R_{24}$  and  $R_{25}$  are each independently H or a  $C_1$ - $C_6$ alkyl,  $C_3$ - $C_7$ cycloalkyl,  $C_1$ - $C_6$ haloalkyl, phenyl,  $C_5$ - $C_7$ cycloheteroalkyl or heteroaryl group each optionally substituted or each of  $R_9$  and  $R_{10}$  or  $R_{12}$  and  $R_{13}$  or  $R_{15}$  and  $R_{16}$  or  $R_{21}$  and  $R_{22}$  or  $R_{24}$  and  $R_{25}$  may be taken together with the nitrogen atom to which they are attached to form a 5- to 7-membered ring optionally containing another heteroatom selected from 10        O, N or S;

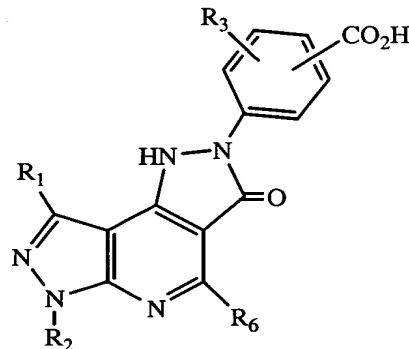
$n$  is 0 or an integer of 1 or 2;

$R_{14}$  is H,  $C_1$ - $C_3$ alkyl or  $C_1$ - $C_3$ haloalkyl;

$R_{19}$  is H or  $C_1$ - $C_3$ alkyl; and

15         $R_{20}$  is a  $C_1$ - $C_6$ alkyl,  $C_3$ - $C_7$ cycloalkyl,  $C_1$ - $C_6$ haloalkyl, phenyl,  $C_5$ - $C_7$ cyclo- heteroalkyl or heteroaryl group each optionally substituted

which process comprises reacting a compound of formula II



(II)

wherein  $R_1$ ,  $R_2$ ,  $R_3$  and  $R_6$  are defined hereinabove with an amine,  $HNR_4R_5$ , in the presence of an activating agent and a solvent.